REPORT DOCUMENTATION PAGE

Form Approved OMB No. 0704-0188

Place reporting ourden for this collection of information is estimated to average 1 hour per response, including the time for reviewing instructions, searching existing data sources, gathering and maintaining the data needed, and completing and reviewing the collection of information. Send comments regarding this burden estimate or any other aspect of this collection of information. Including suggestions for reducing this burden, to Washington Headquarters Services, Directorate for Information Operations and Reports, 1215 Jefferson Casus Highway, Suite 1204, Arington, 74, 22202-4302, and to the Office of Management and Budget, Paperwork Reduction Project (0704-0188), Washington, DC 20503. 3. REPORT TYPE AND DATES COVERED Final 1. AGENCY USE ONLY (Leave plank) | 2. REPORT DATE | Sept 16, 1995 5. FUNDING NUMBERS Neuronal Micronets as Nodal Elements G N00014-92-J-1923 6. AUTHOR(S) Thomas Huntington Brown 8. PERFORMING ORGANIZATION 7. PERFORMING ORGANIZATION NAME(S) AND ADDRESS(ES) REPORT NUMBER Yale University, Grant and Contract Administration, 12 Prospect Place, New Haven, CT 06511-3516 Attn: Ms Sally Tremaine, Associate Director 10. SPONSORING / MONITORING 9. SPONSORING / MONITORING AGENCY NAME(S) AND ADDRESS(ES) AGENCY REPORT NUMBER 11. SUPPLEMENTARY NOTES Attached letter from Thomas H. Brown dated Sept 16, 1996; attached Report of Inventions form. 12b. DISTRIBUTION CODE 12a. DISTRIBUTION / AVAILABILITY STATEMENT No known restrictions on release of information 13. ABSTRACT (Maximum 200 words). We have been working on developing a computationally efficient way to emulate neurons and to emulate circuits and networks of same. We made considerable progress in compressing "realistic" representations of neuronal computations into what we consider functionally equivalent input/output devices, which are now being incorporated into dynamic networks that learn associations and encode time. Our initial hypothesis about how to do this was rejected. Our new hypothesis offers great promise for scaling. This newer hypothesis resulted from examining simulations of "realistic" neurons and thinking about the scaling problem. The latter was funded by the ONR. 15. NUMBER OF PAGES 14. SUBJECT TERMS Artificial neural networks; associative learning; Hebbian 16. PRICE CODE learning; competitive learning; dynamic processing elements. 20. LIMITATION OF ABSTRACT 19. SECURITY CLASSIFICATION SECURITY CLASSIFICATION 17. SECURITY CLASSIFICATION OF ABSTRACT OF THIS PAGE OF REPORT

Yale University

Department of Psychology P.O. Box 11A Yale Station New Haven, Connecticut 06520-7447 Campus address: 2 Hillhouse Avenue.

September 16, 1995

Defense Technical Information Center Building 5, Cameron Station Alexandria, VA 22304-6145

Re: Final Technial Report on ONR grant N00014-92-1923

19960619 006

Dear Sirs/Mesdames:

This is the final technical report on grant N00014-92-J-1923. This information has also been sent to Dr. Thomas McKenna, the Administrative Grants Officer, and the Director of the Naval Research Laboratory.

The overarching goal of this grant "Neuronal Micronets as Nodal Elements" (N00014-92-J-1923) application was to determine the computational significance of the amount and type of information processing that actually occurs in the nervous system at the single-neuron level.

The <u>basic thesis</u> was that a neuron can be compared to a multi-layered artificial neural network (ANN). The question is, what kind of ANN best captures neuronal computations? This question raises several others: How do neurons differ from the processing elements (PEs) used in connectionistic studies? What do single neurons compute? We have been addressing the latter question through compartmental simulations of hippocampal neurons containing Hebbian synapses.

I used the term "micronet" to refer to the type of ANN that can capture neuronal computations¹. I concluded that micronets are deep (many layers of PEs) but narrow (relatively few PEs per layer). To function like a neuron, the PEs must operate continuously and asynchronously. There is no clock. Time is its own representation. Connections within a micronet are assumed not to be modifiable, but connections among micronets can exhibit use-dependent modifications, which can be Hebbian. The PE activation function has a passive memory that decays rapidly and exponentially as a function of time.

I requested an AASERT for Sean Murphy, a neuroscience graduate student interested in this problem, but this application was not funded. Murphy went on however, working with another professor in my department, to begin building ANNs that could emulate the input-output functions of neurons, and this grew into part of his dissertation, which was just completed². As I had anticipated, Murphy concluded that an appropriate neural network can indeed emulate a relatively realistic neuronal model in terms of gross input-output functions.

In working with one of my former students, we reached a similar conclusion, based on less formal or extensive analysis. The main difference, however, was that we concluded that, once learning is involved, the circumstance is quite different. And learning was an essential part of the problem we sought to understand. What follows summarizes some of the reasons for rejecting our initial basic thesis that a neuron can be compared to a multilayered ANN. The point is not that the enterprise cannot be done in principle, but that this approach is much too cumbersome relative to certain alternatives, if one is interested in learning.

Part of the reason is that Hebbian learning depends on the electrotonic structure of the cells and this is not easy to capture in connectionist models³⁻⁹. Hebbian learning depends on the amplitude of the signal at the site of the synaptic input. This in turn means that voltage transfers to and from that site to and from every other synapse must be modeled. In addition, the effects of spiking in the soma must be represented in terms of voltage transfer back through the dendritic tree to the synapses.

These voltage transfers are in general not symmetrical—the attenuation from point i to point j is not the same as from j to i and they are frequency dependent i0. Furthermore, nonlinear membrane responses, which our modeling suggested to be important i1-13, particularly the backpropagation of action potentials into the dendrites, are very difficult to incorporate in an ANN that includes Hebbian learning but have recently been suggested based on experimental data.

Why this insistence on Hebbian learning? Based on first principles, it has always been clear that Hebbian synapses were theoretically important^{4,13,14} and we and others had previously shown them to exist in hippocampus. Now it seems that Hebbian synapses are extremely widespread. One sees such synapses in brain regions other than the hippocampus. One even sees such synapses in lower vertebrates. Most recently, it seems that they may even exist in invertebrates.

Thus we continued to explore electrotonic structure in order to understand better how it might interact with active membrane and Hebbian learning 16-20. At the same time, we developed advanced methods to gain a better understanding of the diversity of cell types and their characteristic active membrane properties 13,21.

My conclusion from this work was that it is computationally easier to use analog models or devices instead of trying to make an ANN emulate what such circuitry would do. If learning can be done off line, or if learning is not involved, then the original idea of representing a neuron as a micronet still makes sense². But from what we now know about the neurophysiology, the nervous system is continuously self-organizing and exhibiting various forms of learning and that these will ultimately depend critically on the electrotonic structure and nonlinear dynamics. Therefore, I focused on the latter. We have gained deep insights into single neuron computations from this enterprise.

We have now begun to formalize the manner in which Hebbian self-organization depends on electrotonic structure 10,22. The effect of nonlinear membrane probably will differ in different classes of cells and for different types of synaptic inputs. What we are learning is that there is no canonical neuron, even within highly circumscribed brain regions. The number of different types is large and we suspect that this means that brain-style computations require a correspondingly large number of elemental devices—in contrast to the conventional assumptions in ANNs.

You will recall that this application was submitted in March of 1992, and I received notice that it would receive no further funding on August of 1993. At this time I requested a no-cost extension.

We are still publishing work from this period and the enterprise motivated us to collect experimental data that confirmed the conclusions based on the simulations mentioned above. Additional manuscripts are in preparation. I am currently working on what I think may be a <u>fundamental new mechanism for how the nervous system encodes time using Hebbian synapses</u>. I am also continuing to work on the design of a simple device that can, with small quantitative variations, learn and self-organize the way real neurons do. In contrast to Murphy's approach², <u>this elemental device would not be an ANN that emulates a particular neuron</u>. Rather I am looking for an analog device that can capture the essence of our insights based on the above considerations. I hope to explore these ideas further through interactions with certain electrical engineers at Yale.

- 1. Brown, T.H., Zador, A.M., Mainen, Z.F., and Claiborne, B.J. Hebbian modifications in hippocampal neurons. In: Long-term potentiation: A debate of current issues, J. Davis and M. Baudry (eds.), pp. 357-389, MIT Press, 1991.
- 2. Murphy, S. D. Temporal dynamics of biological neural network models. Doctoral Dissertation. Yale University. 1995.
- 3. Brown, T.H., Zador, A.M., Mainen, Z.F., and Claiborne, B.J. Hebbian computations in hippocampal dendrites and spines. In: *Single Neuron Computation*, T. McKenna, J. Davis, and S.F. Zornetzer (eds.), pp. 81-116, Academic Press, 1992.
- 4. Brown, T.H., Kairiss, E.W., and Keenan, C.L. Hebbian synapses—biophysical mechanisms and algorithms. *Ann. Rev. Neurosci.*, 13:475-511, 1990.
- 5. Zador, A.M., Koch, C., and Brown, T. H. Biophysical model of a hebbian synapse, *Proc. Nat. Acad Sci.*, 6718-6722, 1990.
- 6. Kairiss, E.W., Mainen, Z.F., Claiborne, B.J., and Brown, T.H. Dendritic control of Hebbian computations. In: *Analysis and Modeling of Neural Systems*, F. Beckman (ed.), pp. 69-83, Kluwer Academic Publishers, 1991.
- 7. Brown, T.H., Zador, A.M., Mainen, Z.F., Claiborne, B.J. Self-organization of Hebbian synapses in hippocampal neurons. In: Advances in Neural Information Processing Systems 3, R. Lippmann, J. Moody, D. Touretzky (eds.), pp. 39-45, Morgan Kaufmann Publishers, 1991.
- 8. Claiborne, B.J., Zador, A.M., Mainen, Z.F., and Brown, T.H. Computational models of hippocampal neurons. In: *Single Neuron Computation*, T. McKenna, J. Davis, and S.F. Zornetzer (eds.), pp. 61-79, Academic Press, 1992.
- 9. Tsai, K.Y., Carnevale, N.T., Brown, T.H. Hebbian learning is jointly controlled by electrotonic and input structure. *Network: Computation in Neural Systems*, 5:1-19, 1994.
- 10. Tsai, K., Carnevale, N. T. and Brown. Efficient mapping from neuroanatomical to electronic space. *Network: Computation in Neural Systems*, 5:21-46, 1994.
- 11. Zador, A.M., Claiborne, B.J., and Brown, T.H. Nonlinear pattern separation in single hippocampal neurons with active dendritic membrane. In: *Advances in Neural Information Processing Systems 4*, J.E. Moody, S.J. Hanson, and R.P. Lippmann (eds.), pp. 51-58, Morgan Kaufmann, 1992.
- 12. Fisher, S. A., Jaffe, D. B., Claiborne, B. J., and Brown, T. H. (1993) Self-organization of Hebbian synapses on a biophysical model of a hippocampal neuron. *Soc Neurosci Abstr* 19: 808.
- 13. Jaffe, D.B., Fisher, S.A., Brown, T.H. Confocal laser scanning microscopy reveals voltage-gated calcium signals within hippocampal dendritic spines. *J. Neurobiol.*, 25(3):220-233, 1994.
- 14. Brown, T. H., and Chattarji, S. Hebbian synaptic plasticity: evolution of the contemporary concept. *Models of Neural Networks II*, E. Domany, J.L. Van Hemmen, and K. Schulten (eds.), pp. 287-314, Springer-Verlag, New York, 1994.
- 15. Brown, T.H., Chattarji, S. Hebbian plasticity. In: The *Handbook of Brain Theory and Neural Networks*, M.A. Arbib (ed.), pp. 454-459, MIT Press, 1995.
- 16. Carnevale, N.T., Tsai, K.Y., Claiborne, B.J. and Brown, T.H. The electrotonic transformation: a tool for relating neuronal form to function. In: Advances in Neural Information Processing Systems 7, Tesauro, G., Touretzky, D.S. and Alspector, J. (eds.), pp. 69-76, Morgan Kaufmann, San Mateo, CA, 1995.
- 17. Carnevale, N.T., Tsai, K.Y., Claiborne, B.J. and Brown, T.H. Qualitative electrotonic comparison of three classes of hippocampal neurons in the rat. In: *The Neurobiology of Computation*, J. Bower (ed.), pp. 67-72, J., Kluwer, Boston, MA, 1995.

- 18. O'Boyle, M.P., Carnevale, N.T., Claiborne, B.J., and Brown, T.H. Sensitivity of hippocampal granule cell models to variations in dendritic diameter. Submitted.
- 19. Carnevale, N.T., Tsai, K.Y. and Brown, T.H. Hebbian self-organization predicted in detailed electrotonic models of hippocampal neurons. In preparation.
- 20. Carnevale, N.T., Tsai, K.Y., Claiborne, B.J., and Brown, T.H. Comparative electrotonic analysis of three classes of rat hippocampal neurons. In preparation.
- 21. Faulkner, B. and Brown, T. (1995) Physiology and morphology of perirhinal cortical neurons. Soc Neurosci Abstr 21: 596.
- 22. Carnevale, N., Tsai, K., and Brown, T. (1995) Prediction of hebbian learning in cells with biologically realistic electrotonic architecture. *Soc Neurosci Abstr* 21: 604.

Sincerely,

Thomas H. Brown, Ph. D Professor of Psychology

Man HD

Professor of Cellular and Molecular Physiology

REPORT OF INVENTIONS AND SUBCONTRACTS

COMB No. 0704-181

(Punuant to "Patent Rights" Contract Gause) (See Instructions on Reverse Side.)

Public reparting burden for this collection of information is estimated to evertage 5 mentions, including the firme for reviewing militarizing and commending and mentioning the data mentioning the collection of information, including suggestion for reducing this burden, to Waisington Manageria Servers, Directorate for information of research frequency factorism to mentioning the collection of the collecti

		ACCOUNTS OF CALLESTING TO SELECT TO	1 - Charles Ct Militable	
IL MAME OF CONTRACTOR/SUBCOMPRACTOR	S CONTINUES MUMBER	24. NAMES OF GOVERNMENT PROME CONTINUED		-
Yale University	1000014-41-2-1463			_1
b. ADDRESS linguide ZIF Codes	A AWARD GATE (TYMMGD)	b. ADORESS (Grande IIP Cost)	4. AWARD DATE (PTMMDD)	4 REPORTING PERIOD (TYLANDO)
12 Prospect Place	7 - 0			10/10/25 MONT
May Laven, CT 06511	10110176			b. 10 95 106/30
A		SECTION 1 - SUBJECT INVENTIONS		
S. SUBJECT INVENTIONS" REQUIRED TO BE BEPORTED BY CONTRACTOR/SURCONTRACTOR	TES EY CONTRACTOR/SURCENTRACTOR OF THE	bret." sa sjatel		
MAME(S) OF INVENTORIS!	TITLE OF INVENTIONS!	C OKCLDSTURE MO.	G. ELECTION TO FILE O. FATENT APPLICATIONS FROM	C. COMMANDER OR INSTRUMENT OR ASSERBENT FORWARDED
(Last. First.Ad)			(a) Ter (b) No (s) Tes (b) No	19 CONTAACTING OFFICIAL
	1/10	JWE		
I EMM OVER OF INVENTORIS NOT EMPLOYED BY CONTRACTOR SUBCONSPACTOR	CONTRACTOR SUBCONINACTOR	4. LIECTTO FORLIGH COUNTINES IN WHICH A PATENT	WHICH A PARENT APPLICATION WILL BE FUED	
(ון (פו אוכשה כו מאהטוכן ווישן קוננו און	Littles Name of Smertar (Lart, Furt, MI)	(1) The of Invention	8 17	(1) foreign Countines of Patent Application
(D) Hame Of Embloyer	(b) Name of Crimoger			and the second
(c) Apprets of Employer linguist Life Cook)	(i) Address of Engacyer (Persone 22P Goos)	786		
	SECTION 11 -	. SUBCONTRACTS (Containing a "Patent Rights" dausel	' dausel	V
4. SUBCONTRACTS AWARDED BY CONTRACTOR/SUBCONTRACTOR IN THEM. TO ITREE	USCOMPLACTOR (IF "Nove," to rester			
HAME OF SURCONTRACTORGS	ASONESS (PROLATE IT!! CORE)	SUBCONTRACT HO.(3) (1) Chave (7) Orto Humber (17444)	DESCHETION OF WORK TO BE MINGORINED UNDER SURCONTRACTISS	1. SUBCONTRACT DATES IFTAMODI (1) Award (2) E Con
	7	JAN!	•	
		SECTION III - CERTIFICATION		*
1. CERTIFICATION OF REPORT BY CONTRACTORYSURCONFRACTOR	2 SURCONTRACTOR (MG	X LEGILLER OF Smest Gariness OF	Mon-mail organization; (X asermanere ode)	
L NAME OF AUTHORIZED COMPACTORY SURCONTRACTOR OFFICIAL THE PIEL MILL MILL POLINEY. SUZZENDE IS.		I certify that the reporting party has procedures for prompt identification and timely disclosure of "Subject Inventions," that such procedures have been followed and that all "Subject inventions" have been reported	ures for prampt identification and tin I followed and that all "Subject inves	nely disclosure of "Subject ntions" have been reported.
b une Director	vi	None of the state		E. DATE SIGNED
Contract Administration	Jministration	Ouscond 184		10/24/95

preylous editions are absolete.

4 40 Eprim 882, OCT 89